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[54] ENDOTHELIAL ENVELOPMENT DRUG **CARRIERS**

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[56]

References Cited

U.S. PATENT DOCUMENTS

		Michaels
4,432,802	2/1984	Harata et al 424/488 X
4,568,536	2/1986	Kronenthal et al 514/965 X
4,624,846	11/1986	Goldenberg 530/387 X
4,671,958	6/1987	Rodwell et al 514/6 X

FOREIGN PATENT DOCUMENTS

1168150 5/1984 Canada. 0087786 9/1983 European Pat. Off. . 0167825 1/1986 European Pat. Off. . 0240424 10/1987 European Pat. Off. . WO83/03426 10/1983 PCT Int'l Appl. . WO84/00294 2/1984 PCT Int'l Appl. . 1516348 7/1978 United Kingdom .

2041517 9/1980 United Kingdom . 2185397 7/1987 United Kingdom .

OTHER PUBLICATIONS

European Search Report, 10/14/88.

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ABSTRACT

This application describes the preparation and in vivo testing of surface coatings and matrix materials, which when applied to or caused to comprise the carriers for drugs and diagnostic agents, and administered in a fashion that allows efficient vascular access, causes the carriers to recognize determinants present or normal or focally diseased endothelium, and induces the following in vivo effects: (1) rapid, partial or total endothelial envelopment of the drug (diagnostic) carrier; (2) sequestration of the carrier and protecting entrapped agent from blood vascular clearance at an early time (2 minutes) when the endothelial pocket which envelops the carrier still invaginates into the vascular compartment; (3) acceleration of the carrier's transport across or through the vascular endothelium and/or subendothelial structures into the tissue compartment (interstitium); and (4) improvement of the efficiency with which the drug (or diagnostic) carrier migrates across the endothelium, or epi-endothelial or subendothelial barriers. such that a lower total drug dose is required to obtain the desired effect relative to that required for standard agents.

4 Claims, 6 Drawing Sheets